	FILE	'CAPLU	JS '	E	NTER	ED AT 07:59:30 ON 16 MAY 2002	
L1		983	s	PRO	TIMAC	HOCYANIDIN	
L2		1	S	L1	AND	RIBOSYLATION (W) INHIBITOR	
L3		209	S	L1	AND	PLANT	
L4		0	S	L3	AND	COMPOSITION	
L5		9	S	L3	AND	COMPOSITION	

- L5 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 2001:766895 CAPLUS
- DN 136:52981
- TI Composition of Grape Skin Proanthocyanidins at Different Stages of Berry Development
- AU Kennedy, James A.; Hayasaka, Yoji; Vidal, Stephane; Waters, Elizabeth J.; Jones, Graham P.
- CS Department of Horticulture Viticulture and Oenology, Adelaide University, Glen Osmond, 5064, Australia
- SO Journal of Agricultural and Food Chemistry (2001), 49(11), 5348-5355 CODEN: JAFCAU; ISSN: 0021-8561
- PB American Chemical Society
- DT Journal
- LA English
- RE.CNT 44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI Composition of Grape Skin Proanthocyanidins at Different Stages of Berry Development
- The compn. of grape (Vitis vinifera L. cv. Shiraz) skin proanthocyanidins was detd. at different stages of berry development. Beginning approx. 3 wk after fruit set and concluding at com. ripeness, the compn. of isolated skin proanthocyanidins was detd. using the following anal. techniques: elemental anal., UV-Vis absorption spectroscopy, reversed-phase HPLC after acid-catalysis in the presence of excess phloroglucinol, gel permeation chromatog., electrospray ionization mass spectrometry (ESI-MS), and 13C NMR. On the basis of these analyses, berry development was correlated with an increase in proanthocyanidin mean d.p., an increase in the proportion of (-)-epigallocatechin extension subunits, and increases in the level of anthocyanins assocd. with the proanthocyanidin fraction. Addnl., data acquired from ESI-MS of the isolates following acid-catalysis in the presence of excess phloroglucinol is consistent with pectin-bound proanthocyanidins.
- ST grape skin proanthocyanidin compn berry development
- IT Color

Grape

Growth and development, plant

(compn. of grape skin proanthocyanidins at different stages of berry development)

IT Growth and development, plant

(fruit ripening; compn. of grape skin proanthocyanidins at different stages of berry development)

- L5 ANSWER 2 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 2001:706541 CAPLUS
- DN 136:246718
- TI Proanthocyanidin composition of red Vitis vinifera varieties from the Douro Valley during ripening: Influence of cultivation altitude
- AU Mateus, Nuno; Marques, Sara; Goncalves, Ana C.; Machado, Jose M.; De Freitas, Victor
- CS Departamento de Qulmica do Porto, Centro de Investigação em Qulmica, Oporto, 4169-007, Port.
- SO American Journal of Enology and Viticulture (2001), 52(2), 115-121 CODEN: AJEVAC; ISSN: 0002-9254
- PB American Society for Enology and Viticulture
- DT Journal
- LA English
- RE.CNT 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI **Proanthocyanidin composition** of red Vitis vinifera varieties from the Douro Valley during ripening: Influence of cultivation altitude
- AB The effect of altitude and its related climatic conditions on the

proanthocyanidin compn. of Touriga Nacional and Touriga Francesa varieties during berry maturation is reported for the 1997 vintage. berry maturation, low altitude is shown to be an important factor favoring the biosynthesis of higher concns. of grape-skin catechin monomers ((+)-catechin, (-)-epicatechin, (-)-epicatechin gallate), procyanidin dimers, trimer C1, as well as total extractable proanthocyanidins. The grapes (skin and seeds) of Touriga Nacional were richer in low mol. wt. flavan-3-ol compds., while Touriga Francesa contained higher concns. of total extractable proanthocyanidins. At harvest, grape-skin dimer content was comprised almost entirely of dimer B1, followed by dimers B2 and B3, whereas C4-C8 linked dimers (B1 to B4) and B2-gallate were the most abundant found in seeds. Dimer B2, which was one of the less important dimers at the early stage of development in seeds, showed a tendency to increase during ripening, while its resp. gallate ester (B2-gallate) markedly decreased. proanthocyanidin compn red Vitis fruit ripening Growth and development, plant (fruit ripening; proanthocyanidin compn. of red Vitis vinifera varieties from Douro Valley during ripening) Grape (proanthocyanidin compn. of red Vitis vinifera varieties from Douro Valley during ripening) Phenols, biological studies Proanthocyanidins RL: BSU (Biological study, unclassified); NPO (Natural product occurrence); BIOL (Biological study); OCCU (Occurrence) (proanthocyanidin compn. of red Vitis vinifera varieties from Douro Valley during ripening) 154-23-4, (+)-Catechin 490-46-0, (-)-Epicatechin 1481-83-0D, Flavan-3-ol, derivs. 12798-57-1, Procyanidin B5 12798-58-2, Procyanidin B6 12798-59-3, Procyanidin B7 12798-60-6, Procyanidin B8 20315-25-7, Procyanidin B1 23567-23-9, Procyanidin B3 29106-49-8, Procyanidin B2 29106-51-2, Procyanidin B4 RL: BSU (Biological study, unclassified); NPO (Natural product occurrence); BIOL (Biological study); OCCU (Occurrence) (proanthocyanidin compn. of red Vitis vinifera varieties from Douro Valley during ripening) ANSWER 3 OF 9 CAPLUS COPYRIGHT 2002 ACS 2000:553206 CAPLUS 133:155161 Cosmetic composition for protecting the scalp from free radicals Herrling, Thomas; Groth, Norbert; Golz-Berner, Karin; Zastrow, Leonhard Coty B. V., Neth. Eur. Pat. Appl., 7 pp. CODEN: EPXXDW Patent German ICM A61K007-40 ICS A61K007-48 FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE ---- --------------EP 1025835 A2 20000809 A3 20010801 EP 2000-250030 20000131 EP 1025835 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO DE 19905127 20000810 DE 1999-19905127 19990201 A1 PRAI DE 1999-19905127 A 19990201 Cosmetic composition for protecting the scalp from free radicals The title compn. comprises an aq. dispersion, emulsion, or hydrogel contg. 0.5-30 wt.% enzymic radical scavenger and 0.1-20 wt.% water-sol. or

-dispersible film-forming agent (shellac and/or dextrin). Thus, a radical

scavenger complex comprised phospholipids 7, quebracho ext. (contg.

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proanthocyanidin oligomers and gallic acid) 2, silkworm ext. (contg. cecropin, amino acids, and vitamins) 1, acerola (Malpighia punicifolia) fruit ext. 1, vitamin C 0.5, and vitamin A 0.5% in a gel base contg. Carbomer, EtOH, and glycerin. This complex 30.0, .alpha.-dextrin 5.0, .beta.-dextrin 2.5, .gamma.-dextrin 5.0, preservative 0.5, and H2O to 100 wt.% were combined to produce a scalp spray.

IT Plant (Embryophyta)

Yeast

(radical scavengers from; cosmetic compn. for protecting the scalp from free radicals)

- L5 ANSWER 4 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 1998:268176 CAPLUS
- DN 129:80945
- TI Chemical composition, rumen degradation, and gas production characteristics of some multipurpose fodder trees and shrubs during wet and dry seasons in the humid tropics
- AU Larbi, A.; Smith, J. W.; Kurdi, I. O.; Adekunle, I. O.; Raji, A. M.; Ladipo, D. O.
- CS Humid Zone Programme, International Livestock Research Institute (ILRI), Ibadan, Nigeria
- SO Animal Feed Science and Technology (1998), 72(1-2), 81-96 CODEN: AFSTDH; ISSN: 0377-8401
- PB Elsevier Science B.V.
- DT Journal
- LA English
- TI Chemical composition, rumen degradation, and gas production characteristics of some multipurpose fodder trees and shrubs during wet and dry seasons in the humid tropics
- AB Seasonal variations in chem. compn., dry matter (DM) and nitrogen (N) degrdn., and gas prodn. characteristics of 18 multipurpose trees and shrubs (MPTs) from the humid lowlands of West Africa were evaluated. MPTs have potential for the development of integrated crop and livestock agroforestry technologies in the region. The expt. was conducted in Ibadan, southwestern Nigeria during the main-wet (Apr.-August) and dry (Dec.-Mar.) seasons. The MPTs were ranked by their degrdn. and gas prodn. characteristics, and these were found to be related to chem. compn. There were wide variations among MPTs in crude protein (CP), neutral detergent fiber (NDF), acid detergent fiber (ADF) and proanthocyanidin contents, DM and N degrdn., and gas prodn. characteristics. Dry matter degrdn. during the dry season ranged from 416 to 868 g kg-1 and for N 508 to 950 g kg-1. Crude protein, and rates of DM and N degrdn. were significantly correlated (r=0.48, P=0.037 for DM and r=0.56, P=0.032 for N). The rates and extents of DM and N degrdn. were significantly correlated with NDF and ADF during the wet season (r=-0.47 to -0.63). The vol. of gas produced (r=-0.48 to -0.67) and initial gas prodn. (r=-0.64 to -0.73) were highly correlated with the NDF and ADF in both seasons. The rate of DM degrdn. was significantly correlated with gas prodn. variables in the minor-wet season. Ranking of the MPTs based on extent of DM and N degrdn., and vol. of gas produced for the main-wet and dry seasons were highly correlated. Based on degrdn. and gas prodn. characteristics in the main-wet and the dry seasons, F. exasperata, S. nodosa, S. siamea, S. spectabilis, G. sepium, L. leucocephala and L. diversifolia were superior in quality to M. thonningii, A. angustissima and P. pterocarpum.

IT Dietary fiber Forage

Plant (Embryophyta)

Tree

(chem. compn., rumen degrdn., and gas prodn. characteristics of some multipurpose fodder trees and shrubs during wet and dry seasons in the humid tropics)

L5 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2002 ACS AN 1996:713825 CAPLUS

- DN 126:17935
- TI Determination of the **composition** of commercial tannin extracts by liquid secondary ion mass spectrometry (LSIMS)
- AU Vivas, Nicolas; Bourgeois, Guy; Vitry, Christiane; Glories, Yves; de Freitas, Victor
- CS Faculte d'OEnologie, Univ. Victor Segalen, Talence, 33405, Fr.
- SO J. Sci. Food Agric. (1996), 72(3), 309-317 CODEN: JSFAAE; ISSN: 0022-5142
- PB Wiley
- DT Journal
- LA English
- TI Determination of the **composition** of commercial tannin extracts by liquid secondary ion mass spectrometry (LSIMS)
- AB The compns. of various com. tannin exts. were detd. by liq. secondary ion mass spectrometry (LSIMS). Spectra were obtained directly from tannin exts. without any pre-sepn. Eight different tannin powders were analyzed: three gallotannins (Chinese, Turkish, tara), three ellagitannins (sweet chestnut, pendunculata oak, sessile oak), one mixed hydrolysable tannin (myrabolans) and one proanthocyanidin (grape seeds). This method enabled the main mols. in these powders to be identified.
- ST tannin detn **plant** source liq SIMS; secondary ion mass spectrometry tannin
- L5 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 1994:187333 CAPLUS
- DN 120:187333
- TI Developmental changes in the concentration and **composition** of flavonoids in skin of a red and a green apple cultivar
- AU Lister, Carolyn E.; Lancaster, Jane E.; Sutton, Kevin H.; Walker, John R. L.
- CS Plant Improv. Div., N Z Inst. Crop and Food Res. Ltd., Christchurch, N. Z.
- SO J. Sci. Food Agric. (1994), 64(2), 155-61 CODEN: JSFAAE; ISSN: 0022-5142
- DT Journal
- LA English
- TI Developmental changes in the concentration and **composition** of flavonoids in skin of a red and a green apple cultivar
- Flavonoids from the skin of Granny Smith, a green apple cultivar, and AB Splendour, a red apple cultivar, were quantified by high-performance liq. chromatog. for two seasons (1989-1990 and 1990-1991). Both cultivars contained a similar compn. and concn. of quercetin glycosides and proanthocyanidins. Splendour also synthesized cyanidin glycosides during ripening. Quercetin glycosides and proanthocyanidins were highest in the skin of very young fruit of Granny Smith and decreased by 50% during fruit development. In Splendour, concns. of quercetin glycosides and proanthocyanidins in the skin decreased by 50% from early to mid-season but then increased during ripening. Cyanidin glycosides in Splendour increased to about 1 mg g-1 fresh wt. during ripening. There were significant differences between the two cultivars but not between years. Total amt. of flavonoids increased throughout the season as fruit surface area increased. For Granny Smith there was an estd. net synthesis per apple of 0.16 mg day-1 quercetin glycosides, 0.1 mg day-1 proanthocyanidins and for Splendour a net synthesis per apple of 0.28 mg day-1 quercetin glycosides, 0.21 mg day-1 proanthocyanidins and during ripening 0.21 mg day-1 cyanidin glycosides. Relative proportions of major quercetin glycosides and proanthocyanidins were stable during fruit development. For Splendour, however, cyanidin glycoside synthesis was accompanied by a corresponding increase in quercetin glycoside and proanthocyanidin synthesis. The data suggest a coordinate regulation of enzymes in the flavonoid biosynthetic pathway during fruit development.
- IT **Plant** growth and development (fruit-ripening, flavonoids of red and green apple cultivars during)

- L5 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 1993:445347 CAPLUS
- DN 119:45347
- TI Developmental changes in the **composition** of proanthocyanidins from leaves of sainfoin (Onobrychis viciifolia Scop.) as determined by HPLC analysis
- AU Koupai-Abyazani, Mohammed R.; McCallum, John; Muir, Alister D.; Bohm, Bruce A.; Towers, G. H. N.; Gruber, Margaret Y.
- CS Dep. Bot., Univ. British Columbia, Vancouver, BC, V6T 1Z4, Can.
- SO J. Agric. Food Chem. (1993), 41(7), 1066-70 CODEN: JAFCAU; ISSN: 0021-8561
- DT Journal
- LA English
- TI Developmental changes in the **composition** of proanthocyanidins from leaves of sainfoin (Onobrychis viciifolia Scop.) as determined by HPLC analysis
- AB Proanthocyanidin (PA) polymer (condensed tannins) were extd. from sainfoin leaves (O. vicifolia) at different stages of plant development. Anal. of the phloroglucinol degrdn. products by HPLC showed that catechin, epicatechin, gallocatechin, and epigallocatechin were present as terminal units at all stages, while gallocatechin and epigallocatechin were the predominant extension units with lesser amts. of epicatechin incorporated at early stages. Catechin was not incorporated as an extension unit. The no.-av. mol. wt. and d.p. increased with leaf development. There was a very distinct change in the isomerization and degree of hydroxylation of the polymer constituents with development. The compn. of cis-isomers decreased from 83 to 48% and the proportion of trihydroxylated B-rings increased from 60 to 90% with increasing leaf maturity.
- ST sainfoin proanthocyanidin development
- IT Plant growth and development
 - (proanthocyanidins of sainfoin in relation to)
- L5 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2002 ACS
- AN 1988:19211 CAPLUS
- DN 108:19211
- TI Chemical composition of barley varieties with different nutrient supplies. III. Concentration of tannins and .beta.-glucans in two-year experiments
- AU Truelsen, Ebbe
- CS Res. Cent. Agric., Dan. Res. Serv. Plant Soil Sci., Lyngby, DK-2800, Den.
- SO Tidsskr. Planteavl (1987), 91(1), 69-76 CODEN: TPLAAV; ISSN: 0040-7135
- DT Journal
- LA English
- TI Chemical composition of barley varieties with different nutrient supplies. III. Concentration of tannins and .beta.-glucans in two-year experiments
- AB Twenty barley varieties were grown in 1982 and 1983, and 21 varieties, including a breeding line Ca 700202, were grown in 1984 and 1985 in pots with increasing N supplies. The concns. of tannins and sol. .beta.-glucans were detd. in the mature grains. The contents of tannins and .beta.-glucans were closely connected to the varieties, therefore highly significant correlations could be calcd. between the two years of expts. Only a low pos. reaction for tannin was found in the proanthocyanidin-free variety Galant, but Cerise, Carina, Ca 700202 and Claret also had fairly low tannin contents. Significant differences between the two years were found in tannin contents. The lowest content of sol. .beta.-glucans was found in the breeding line Ca 700202 and the varieties Triumph, Mandolin and Yriba. No significant differences could be found in the content of .beta.-glucans between the two years. Increasing supplies of N caused increases in the content of sol. .beta.-glucans, whereas only small changes in the content of tannins were found.

- ITPlant breeding and selection (of barley, glucans and tannin content in relation to) L5 ANSWER 9 OF 9 CAPLUS COPYRIGHT 2002 ACS AN 1983:555346 CAPLUS DN 99:155346 Proanthocyanidins of barley and sorghum; composition as a TIfunction of maturity of barley ears ΑU Brandon, Michael J.; Foo, Lai Yeap; Porter, Lawrence J.; Meredith, Peter Dep. Sci. Ind. Res., Wheat Res. Inst., Petone, N. Z. CS SO Phytochemistry (1982), 21(12), 2953-7 CODEN: PYTCAS; ISSN: 0031-9422 DTJournal LΆ English ΤI Proanthocyanidins of barley and sorghum; composition as a function of maturity of barley ears AB Sorghum vulgare Seeds contain a proanthocyanidin polymer consisting largely of 2,3-cis procyanidin units with no.-av. mol. wt. 2500. Hordeum vulgare Ears contain low levels of proanthocyanidin oligomers contg. 2-4 units, and composed largely of 2,3-trans procyanidin and prodelphinidin units, with catechin as the terminal unit. The concn. of the oligomers in barley ears was const. throughout the 33-day growth and ripening period.
- ST proanthocyanidin barley sorghum maturation; tannin condensed barley sorghum

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                  frequency
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                 TRCTHERMO no longer available
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379881 INHIBITOR

401899 INHIBITORS

L1 619425 INHIBITOR

(INHIBITOR OR INHIBITORS)

=> s ll and ribosylation

5143 RIBOSYLATION

30 RIBOSYLATIONS

5146 RIBOSYLATION

(RIBOSYLATION OR RIBOSYLATIONS)

L2 835 L1 AND RIBOSYLATION

=> s 12 and ADP

52454 ADP

65 ADPS

52487 ADP

(ADP OR ADPS)

L3 768 L2 AND ADP

=> s 13 and composition

587900 COMPOSITION

235499 COMPOSITIONS

819660 COMPOSITION

(COMPOSITION OR COMPOSITIONS)

1148107 COMPN

449414 COMPNS

1399752 COMPN

(COMPN OR COMPNS)

1831873 COMPOSITION

(COMPOSITION OR COMPN)

L4 48 L3 AND COMPOSITION

=> s 14 and diptheria

124 DIPTHERIA

L5 0 L4 AND DIPTHERIA

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'=> s 14 and treatment
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         155039 TREATMENTS
        1746876 TREATMENT
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             12 L4 AND TREATMENT
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 => s 16 and diphtheria
           7107 DIPHTHERIA
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          50619 INFECTIONS
         193216 INFECTION
                  (INFECTION OR INFECTIONS)
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           3001 ENTEROTOXINS
          7348 ENTEROTOXIN
                  (ENTEROTOXIN OR ENTEROTOXINS)
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L1
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L2
            835 S L1 AND RIBOSYLATION
L3
            768 S L2 AND ADP
L4
             48 S L3 AND COMPOSITION
L_5
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L6
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L11
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L12
              1 S L4 AND ENTEROTOXIN
=> dis 112 bib abs
L12 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS
   1992:442296 CAPLUS
AN
DN
   117:42296
    Nucleotide regulation of heat-stable enterotoxin receptor
TI
    binding and of guanylate cyclase activation
```

Katwa, Laxmansa C.; Parker, Charlotte D.; Dybing, Jody K.; White, Arnold ΑU

John M. Dalton Res. Cent., Univ. Missouri, Columbia, MO, 65211, USA CS SQ

Biochem. J. (1992), 283(3), 727-35 CODEN: BIJOAK; ISSN: 0306-3275

DT Journal

LA English

Certain nucleotides were found to regulate the binding of the Escherichia AB coli heat-stable enterotoxin (STa) to its receptor in pig intestinal brush border membranes. ATP and adenine nucleotide analogs inhibited 125I-STa binding, while guanine nucleotide analogs stimulated binding, with maximal effects at 0.5-1.0 $\ensuremath{\text{mM}}\xspace$. The strongest inhibitors were adenosine 5'-[.beta..gamma.-imido]triphosphate (App[NH]p) (36%) and adenosine 5'-[.beta.-thio]diphosphate (ADP [S]) (41%). Inhibition did not require Mg2+, and was blocked by p-chloromercuribenzenesulfonate (PCMBS). Stimulation of binding required Mg2+, was not prevented by PCMBS, and was maximal with GDP[S] (41%). While App[NH]p and MgGDP[S] appeared to be acting at different sites, they also interfered with each other. These nucleotides exerted only inhibitory effects on STa-stimulated guanylate cyclase activity, in contrast with the stimulatory effects of adenine nucleotides on atrial natriuretic peptide-stimulated guanylate cyclase. Inhibition by low concns. of MgApp[NH]p and MgATP was weaker above 0.1 mM, while MgGDP[S] and magnesium guanosine 5'-[.gamma.-thiol]triphosphate (MgGTP[S]) inhibited in a single phase. Inhibition by MgApp[NH]p, at all concns., was competitive with the substrate (MgGTP), as was that by MgGDP[S] and MgGTP[S]. Whereas membrane guanylate cyclases usually show pos. co-operative kinetics with respect to the substrate, STa-stimulated activity exhibited Michaelis-Menten kinetics with respect to MgGTP. changed to pos. co-operativity when Lubrol PX was the activator, or when the substrate was MnGTP. These results suggest the presence of both a regulatory and a catalytic nucleotide-binding site, which do not interact co-operatively with STa activation.

=> dis 110 bib abs

L10 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS

ΑN 2001:380730 CAPLUS

DN 134:348261

Fusion proteins that specifically inhibit protein synthesis in neuronal TΙ ΙN

Francis, Jonathan W.; Brown, Robert H., Jr.; Murphy, John R.; Vanderspek, Johanna C.; Oyler, George PΑ

The General Hospital Corporation, USA; Trustees of Boston University; University of Maryland, Baltimore SO

PCT Int. Appl., 35 pp. CODEN: PIXXD2

DT Patent

English

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE WO 2001036588 A2 20010525

PT W: CA, JP, US

WO 2000-US31680 20001116 RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

PT, SE, TR

PRAI US 1999-165807P Ρ 19991116

This invention relates to compns. and methods for the specific inhibition of protein synthesis in neuronal cells leading to neuronal cell death. More specifically, the invention relates to hybrid protein mols. that show high specificity for, and increased cytotoxicity in, neuronal cells. Such hybrid mols. are useful in a variety of conditions where localized inhibition of neuronal cell function is desirable. A fusion gene encoding the first 388 amino acids of diphtheria toxin linked to

tetanus toxin fragment C was constructed, expressed in Escherichia coli strain BL21(DE3) and purified. Following overnight treatment of cultured striatal neurons or N18-RE-105 cells with various concns. of the chimeric toxin, the chimeric toxin was shown to be a potent inhibitor of cellular protein synthesis.

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=> dis 19 1-6 bib abs
      ANSWER 1 OF 6 CAPLUS COPYRIGHT 2002 ACS
 L9
      1999:795640 CAPLUS
 ΑN
 DN
      132:44996
      Wound treatment through inhibition of adenosine diphosphate
 TI
      ribosyl transferase
 ΙN
      Leibovich, Samuel J.
 PA
      University of Medicine and Dentistry of New Jersey, USA
 SO
      PCT Int. Appl., 65 pp.
      CODEN: PIXXD2
 DT
      Patent
 LA
      English
 FAN.CNT 1
      PATENT NO.
                     KIND DATE
                                      APPLICATION NO. DATE
      -----
 PT
      WO 9963982
                      A1 19991216
                                         WO 1999-US13264 19990611
         W: AU, CA, JP
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
      CA 2329160
                       AA . 19991216
                                           CA 1999-2329160 19990611
      AU 9944383
                       A1 19991230
                                           AU 1999-44383
                                                            19990611
                                          EP 1999-927490
     EP 1085859
                      A1
                            20010328
                                                            19990611
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
 PRAI US 1998-88924P
                       Ρ
                            19980611
     WO 1999-US13264
                       W
                            19990611
     A method is disclosed for healing a wound in a mammal which comprises (A)
AB
     providing a therapeutic wound healing compn. comprising a
     therapeutically effective amt. of an inhibitor of mono-
     ADP-ribosyl transferase to inhibit ADP-
     ribosylation of vascular endothelial growth factor, and (B)
     contacting the therapeutic wound healing compn. with a wound in
     a mammal. Also disclosed are wound healing compns. and methods
     for prepg. and using the wound healing compns. and the
     pharmaceutical products in which the therapeutic compns. may be
     used. Further disclosed are therapeutic dermatol.-wound healing
     compns. useful to minimize and treat diaper dermatitis and methods
     for prepg. and using the therapeutic dermatol.-wound healing
     compns.
RE.CNT 1
              THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L9
     ANSWER 2 OF 6 CAPLUS COPYRIGHT 2002 ACS
     1992:4569 CAPLUS
AN
DN
     116:4569
     Determination of G-protein levels, ADP-ribosylation by
TΤ
     cholera and pertussis toxins and the regulation of adenylyl
     cyclase activity in liver plasma membranes from lean and genetically
     diabetic (db/db) mice
ΑU
     Palmer, Timothy M.; Houslay, Miles D.
    Inst. Biochem., Univ. Glasgow, Glasgow, G12 8QQ, UK
CS
SO
    Biochim. Biophys. Acta (1991), 1097(3), 193-204
    CODEN: BBACAQ; ISSN: 0006-3002
ĎΤ
    Journal
LA
    English
    Liver plasma membranes prepd. from genetically diabetic (db/db) mice
AΒ
    expressed levels of Gi .alpha.-2, Gi .alpha.-3 and G-protein
```

.beta.-subunits that were reduced by some 75, 63 and 73% compared with levels seen in membranes from lean animals. In contrast, there were no significant differences in the expression of the 42 and 45 kDa forms of Gs .alpha.-subunits. Pertussis toxin-catalyzed ADPribosylation of membranes from lean animals identified a single 41 kDa band whose labeling was reduced by some 86% in membranes from diabetic animals. Cholera toxin-catalyzed ADP-ribosylation identified two forms of Gs .alpha.-subunits whose labeling was about 4-fold greater in membranes from diabetic animals compared with those from lean animals. Maximal stimulations of adenylyl cyclase activity by forskolin (100 .mu.M), GTP (100 .mu.M), p[NH]ppG (100 .mu.M), NaF (10 mM) and glucagon (10 .mu.M) were similar in membranes from lean and diabetic animals, whereas stimulation by isoprenaline (100 .mu.M) was lower by about 22%. Lower concns. (EC50-60 nM) of p[NH]ppG were needed to activate adenylyl cyclase in membranes from diabetic animals compared to those from lean animals (EC50-158 nM). As well as causing activation, p[NH]ppG was capable of eliciting a pertussis toxin-sensitive inhibitor effect upon forskolin-stimulated adenylyl cyclase activity in membranes from both lean and diabetic animals. However, maximal inhibition of adenylyl cyclase activity in membranes from diabetic animals was reduced to around 60% of that found using membranes from lean animals. Pertussis toxin-treatment in vivo enhanced maximal stimulation of adenylyl cyclase by glucagon, isoprenaline and p[NH]ppG through a process suggested to be mediated by the abolition of functional Gi activity. The lower levels of expression of G-protein .beta.-subunits, in membranes from diabetic compared with lean animals, is suggested to perturb the equil. between holomeric and dissocd. G-protein subunits. It is proposed that this may explain both the enhanced sensitivity of adenylyl cyclase to stimulation by p[NH]ppG in membranes from diabetic animals and the altered ability of pertussis and cholera toxins to catalyze the ADP-ribosylation of G-proteins in membranes from these two animals.

- ANSWER 3 OF 6 CAPLUS COPYRIGHT 2002 ACS L9
- 1991:527584 CAPLUS AN
- DN 115:127584
- Glucocorticoid receptor activation leads to up-regulation of adenosine A1 receptors and down-regulation of adenosine A2 responses in DDT1 MF-2 smooth muscle cells ΑU
- Gerwins, Paer; Fredholm, Bertil B.
- CS Dep. Pharmacol., Karolinska Inst., Stockholm, S-104 01, Swed.
- SO Mol. Pharmacol. (1991), 40(2), 149-55 CODEN: MOPMA3; ISSN: 0026-895X
- DT Journal
- LA English
- The effect of glucocorticoid treatment of DDT1 MF-2 smooth AΒ muscle cells on the signaling via 2 adenosine receptors with opposing actions on cAMP generation was examd. Treatment with dexamethasone caused a dose- and time-dependent increase in the no. of adenosine Al receptors but did not affect the KD or the proportions of receptors in high and low affinity states. The EC50 was 1 $\overline{\text{nM}}$ dexamethasone, and maximal response was achieved after 24 h. The no. of receptors was increased by approx. 50%. Other steroid hormones, including aldosterone, progesterone, testosterone, and estrogen, were much less effective, and addn. of the glucocorticoid receptor antagonist RU 486 or the protein synthesis inhibitor cycloheximide prevented the up-regulation, showing that the effect was mediated via a glucocorticoid receptor-specific mechanism that involves protein synthesis. In dexamethasone-treated cells the Al receptor agonist (-)-N6phenylisopropyladenosine [(R)-PIA] was 3-times more potent as an inhibitor of cAMP formation induced by isoprenaline than in untreated cells. ADP ribosylation of inhibitory GTP-binding proteins by pertussis toxin completely prevented (R)-PIA from inhibiting cAMP accumulation. A further anal. of the different GTP-binding proteins, including the 3 Gi subtypes (Gil, Gi2, and

Gi3), revealed no quant. or qual. change after dexamethasone treatment. In addn., the adenosine A2 receptors were down-regulated, as indicated by the fact that the ability of the A2 receptor agonist 5'-N-ethylcarboxamidoadenosine to increase cAMP formation was decreased by 20-30% in dexamethasone-treated cells. In summary, Al and A2 receptors on the same cell are differentially regulated by glucocorticoids and this has functional importance in the regulation of cAMP accumulation.

- L9 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2002 ACS
- ΑN 1991:119555 CAPLUS
- DN 114:119555
- Mechanism of cytokine inhibition of .beta.-adrenergic agonist stimulation TIof cyclic AMP in rat cardiac myocytes. Impairment of signal transduction
- Chung, Mina K.; Gulick, Tod S.; Rotondo, Russell E.; Schreiner, George F.; AII Lange, Louis G. CS
- Med. Cent., Washington Univ., St. Louis, MO, 63110, USA Circ. Res. (1990), 67(3), 753-63
- SO CODEN: CIRUAL; ISSN: 0009-7330
- DT Journal
- LAEnglish
- Activated immune cells produce a sol. inhibitor of cardiac AB myocyte contractile and cAMP (cAMP) responses to .beta.-adrenergic stimulation. To examine the mechanics of this effect, metabolic assays were conducted on cultured rat cardiac myocytes incubated in the presence and absence of supernatants harvested from rat activated splenocyte cultures. Intracellular cAMP accumulation in response to isoproterenol was inhibited by up to 74% in a dose-dependent fashion by conditioned media contg. sol. cytokines from activated immune cells. By use of myocyte cultures in which contaminating non-myocyte proliferation was inhibited by nonlethal irradn., this phenomenon was shown to be independent of mitogenic effects. Isobutylmethylxanthine, a phosphodiesterase inhibitor, did not ablate cytokine-induced inhibition of cAMP accumulation. Parameters of .beta.-adrenergic receptor binding and affinity were also unaffected. CAMP suppression was maintained after cholera toxin stimulation of cAMP prodn. via stimulatory G protein ADP-ribosylation. CAMP inhibition was not apparent when cells were stimulated with forskolin, a direct adenylate cyclase activator. Importantly, pertussis toxin treatment significantly ablated cytokine-induced cAMP inhibition. Thus, interference with agonist-occupied .beta.-adrenergic receptor coupling to adenylate cyclase to produce cAMP and subsequent contractile responses is induced by a factor(s) elaborated by activated immune cells. This interference occurs at the level of signal transduction across the membrane, can be overridden by pertussis toxin, and may involve changes in the coupling of the stimulatory/inhibitory $\bar{\mathsf{G}}$ proteins to adenylate cyclase. These results demonstrate a novel mechanism of cytokine-induced myocyte dysfunction and may have important pathophysiol. ramifications in immune-mediated myocardial diseases.
- L9 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS
- 1990:491555 CAPLUS
- DN 113:91555
- Pseudomonas exotoxin A prevents .beta.-adrenoceptor-induced up-regulation ΤI of Gi protein .alpha.-subunits and adenylyl cyclase desensitization in rat heart muscle cells ΑU
- Reithmann, Christopher; Gierschik, Peter; Mueller, Ursula; Werdan, Karl; Jakobs, Karl H.
- Pharmakol. Inst., Univ. Heidelberg, Heidelberg, D-6900, Fed. Rep. Ger. CS
- Mol. Pharmacol. (1990), 37(5), 631-8 CODEN: MOPMA3; ISSN: 0026-895X
- DTJournal
- LA English
- Exposure of rat heart muscle cells to noradrenaline (1 .mu.M) for 48 h led AΒ to a decrease in the no. of .beta.1-adrenoceptors of 50% and a concomitant

decrease in adenylyl cyclase stimulation by isoprenaline and forskolin of .apprx.60 and 30%, resp. In addn., the levels of 2 inhibitor guanine nucleotide-binding protein (Gi protein) .alpha.-subunits (Gi.alpha.40 and Gi.alpha.41) were increased in membranes of noradrenaline-treated cells. Evidence is presented that noradrenaline induces this increase by activation of .beta.-adrenoceptors. First, the noradrenaline action was mimicked by the .beta.-adrenoceptor agonist isoprenaline. Second, .beta.-adrenoceptor blockade by timolol but not .alpha.-adrenoceptor blockade by prazosin prevented the noradrenaline-induced up-regulation of Gi.alpha. proteins. Furthermore, timolol but not prazosin abolished the noradrenaline-induced down-regulation of .beta.1-adrenoceptors and the decreases in receptor-dependent (isoprenaline) and -independent (forskolin) adenylyl cyclase stimulation. The specific protein synthesis inhibitor Pseudomonas exotoxin A was used to study whether the noradrenaline-induced up-regulation of Gi .alpha.-subunits depends on increased synthesis of these proteins. This toxin inhibits peptide chain elongation by ADP-ribosylating elongation factor 2. Treatment of rat heart muscle cells with Pseudomonas exotoxin A (1 ng/mL) completely prevented the noradrenaline-induced increase in Gi.alpha. proteins, measured by both pertussis toxin-catalyzed ADPribosylation and immunoblotting with anti-gi.alpha. antibodies. Most importantly, Pseudomonas exotoxin A also completely prevented the noradrenaline-induced decrease in forskolin-stimulated adenylyl cyclase Furthermore, the noradrenaline-induced decrease in isoprenaline-stimulated adenylyl cyclase activity was significantly attenuated by the toxin although the down-regulation of .beta.l-adrenoceptors caused by noradrenaline treatment was not affected. The data presented suggest that prolonged activation of .beta.-adrenoceptors in rat heart muscle cells, in addn. to causing a receptor down-regulation, induced the synthesis of Gi.alpha. proteins, which then apparently mediate a decreased adenylyl cyclase responsiveness. The data, addnl., suggest that the synthesis of Gi.alpha. proteins is under control of the activity of the adenylyl cyclase system and that altered levels of these proteins may play a major role in long term regulation of signal transduction by this enzyme.

- L9 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2002 ACS
- AN 1987:150273 CAPLUS
- DN 106:150273
- Neuropeptide Y inhibits cardiac adenylate cyclase through a pertussis toxin-sensitive G protein
- AU Kassis, Shouki; Olasmaa, Marjut; Terenius, Lars; Fishman, Peter H.
- CS Membrane Biochem. Sect., Natl. Inst. Neurol. Commun. Dis. Stroke, Bethesda, MD, 20892, USA
- SO J. Biol. Chem. (1987), 262(8), 3429-31 CODEN: JBCHA3; ISSN: 0021-9258
- DT Journal
- LA English
- Neuropeptide Y [82785-45-3], a major neuropeptide and potent vasoconstrictor, inhibited (-)-isoproterenol [51-31-0]-stimulated adenylate cyclase [9012-42-4] activity in cultured rat atrial cells as well as in atrial membranes. Prior treatment of the cells with pertussis toxin blocked the inhibitory action of neuropeptide Y. Pertussis toxin is known to uncouple the receptors for other inhibitors of adenylate cyclase by ADP-ribosylation of the .alpha.-subunit of Gi, the inhibitory guanine nucleotide-binding component of adenylate cyclase. The toxin specifically catalyzed the ADP-ribosylation of a 41-kilodalton atrial membrane protein which corresponded to the Gi subunit. Thus, neuropeptide Y may mediate some of its physiol. effects through specific receptors linked to the inhibitory pathway of adenylate cyclase.

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2001:380730 CAPLUS
  AN
  DN
       134:348261
       Fusion proteins that specifically inhibit protein synthesis in neuronal
  TI
       Francis, Jonathan W.; Brown, Robert H., Jr.; Murphy, John R.; Vanderspek,
  ΙN
       Johanna C.; Oyler, George
       The General Hospital Corporation, USA; Trustees of Boston University;
  PA
       University of Maryland, Baltimore
  SO
       PCT Int. Appl., 35 pp.
       CODEN: PIXXD2
  DT
       Patent
 LA
      English
 FAN.CNT 1
      PATENT NO.
                      KIND DATE
                                            APPLICATION NO. DATE
       ______
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 ΡI
      WO 2001036588
                       A2 20010525
                                            WO 2000-US31680 20001116
          W: CA, JP, US
          RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
              PT, SE, TR
 PRAI US 1999-165807P
                       P
                             19991116
      This invention relates to compns. and methods for the specific
      inhibition of protein synthesis in neuronal cells leading to neuronal cell
      death. More specifically, the invention relates to hybrid protein mols.
      that show high specificity for, and increased cytotoxicity in, neuronal
      cells. Such hybrid mols. are useful in a variety of conditions where
      localized inhibition of neuronal cell function is desirable. A fusion
      gene encoding the first 388 amino acids of diphtheria toxin
      linked to tetanus toxin fragment C was constructed, expressed in
      Escherichia coli strain BL21(DE3) and purified. Following overnight
      treatment of cultured striatal neurons or N18-RE-105 cells with
      various concns. of the chimeric toxin, the chimeric toxin was shown to be
      a potent inhibitor of cellular protein synthesis.
 => dis hist
      (FILE 'HOME' ENTERED AT 08:38:59 ON 16 MAY 2002)
     FILE 'CAPLUS' ENTERED AT 08:39:13 ON 16 MAY 2002
I.1
         619425 S INHIBITOR
L2
            835 S L1 AND RIBOSYLATION
L3
            768 S L2 AND ADP
L4
             48 S L3 AND COMPOSITION
L5
              0 S L4 AND DIPTHERIA
L6
             12 S L4 AND TREATMENT
L7
              0 S L5 AND DIPHTHERIA
^{18}
              1 S L6 AND DIPHTHERIA
L9
              6 S L6 AND PERTUSSIS
L10
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L11
              0 S L6 AND INFECTION
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=> s 14 and proanthocyanidin
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                 (PROANTHOCYANIDIN OR PROANTHOCYANIDINS)
L13
             1 L4 AND PROANTHOCYANIDIN
=> s 13 and proanthocyanidin
          983 PROANTHOCYANIDIN
          1799 PROANTHOCYANIDINS
         1920 PROANTHOCYANIDIN
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ANSWER 1 OF 1 CAPLUS COPYRIGHT 2002 ACS

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=> dis 14 bib abs

L4 ANSWER 1 OF 48 CAPLUS COPYRIGHT 2002 ACS

AN 2002:185378 CAPLUS

DN 136:212896

TI Gene markers useful for detecting skin damage in response to ultraviolet radiation

IN Blumenberg, Miroslav

PA New York University School of Medicine, USA

SO PCT Int. Appl., 274 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PΙ

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2002020849 A2 20020314 WO 2001-US28214 20010907

W: AU, CA, JP, SG

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR

PRAI US 2000-231061P P 20000908

The cellular response to UV radiation exposure has been characterized on the mol. level through the use of high d. gene array technol. Nucleic acid mols. and protein mols., the expression of which are repressed or induced in response to UV radiation exposure, are identified according to a temporal pattern of altered expression post UV radiation exposure. Methods are disclosed that utilized these UV radiation-regulated mols. as markers for UV radiation exposure. Other screening methods of the invention are designed for the identification of compds. that modulate the response of a cell to UV radiation exposure. The invention also provides compns. useful for drug screening or pharmaceuticals purposes.

=> dis 13 bib abs

- L3 ANSWER 1 OF 768 CAPLUS COPYRIGHT 2002 ACS
- AN 2002:351551 CAPLUS
- Regulation and recruitment of phosphatidylinositol 4-kinase on immature secretory granules is independent of ADP-ribosylation
- AU Panaretou, Christina; Tooze, Sharon A.
- CS The Secretory Pathways Laboratory, Cancer Research UK London Institute,
 London, WC2A 3PX, UK
- SO Biochemical Journal (2002), 363(2), 289-295 CODEN: BIJOAK; ISSN: 0264-6021
- PB Portland Press Ltd.
- DT Journal
- LA English
- AB Heterotrimeric G-proteins, as well as small GTPases of the Rho and ADP-ribosylation factor (ARF) family, are implicated in the regulation of lipid kinases, including PtdIns 4-kinases and PtdIns(4)P 5-kinases. Here, we describe a PtdIns 4-kinase activity on immature secretory granules (ISGs), regulated secretory organelles formed from the trans-Golgi network (TGN), and investigate the regulation of PtdIns4P levels on these membranes. Over 50% of the PtdIns 4-kinase activity on ISGs is inhibited by both a low concn. of adenosine and the monoclonal antibody 4C5G, a specific inhibitor of the type II PtdIns 4-kinase. Treatment of ISGs with mastoparan 7 (M7) stimulates the type II PtdIns 4-kinase via pertussis-toxin-sensitive Gi/GO proteins, which, in contrast with previous results obtained with chromaffin granules, does not require Rho A, B or C. M7 treatment also leads to an inhibition in the recruitment of ARF to ISG membranes: this inhibition is not dependent on

Gi/GO activation, and is not linked to the stimulation of PtdIns 4-kinase obsd. with M7. PtdIns 4-kinase activity on ISGs is not regulated by myristoylated ARF1-GTP, in contrast with results obtained with Golgi membranes, whereas ARF1-GTP does regulate the prodn. of PtdIns(4,5)P2. Our results suggest that the regulation of PtdIns 4-kinase on the ISGs differs in comparison with that on the TGN, and might be related to a specific requirement of ISG maturation.

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FILE 'MEDLINE' ENTERED AT 08:52:37 ON 16 MAY 2002

FILE LAST UPDATED: 15 MAY 2002 (20020515/UP). FILE COVERS 1958 TO DATE.

On April 22, 2001, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE now contains IN-PROCESS records. See HELP CONTENT for details.

MEDLINE is now updated 4 times per week. A new current-awareness alert frequency (EVERYUPDATE) is available. See HELP UPDATE for more information.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2001 vocabulary. Enter HELP THESAURUS for details.

The OLDMEDLINE file segment now contains data from 1958 through 1965. Enter HELP CONTENT for details.

Left, right, and simultaneous left and right truncation are available in the Basic Index. See HELP SFIELDS for details.

THIS FILE CONTAINS CAS REGISTRY NUMBERS FOR EASY AND ACCURATE SUBSTANCE IDENTIFICATION.

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(FILE 'HOME' ENTERED AT 08:38:59 ON 16 MAY 2002)

FILE 'CAPLUS' ENTERED AT 08:39:13 ON 16 MAY 2002 L1619425 S INHIBITOR L2835 S L1 AND RIBOSYLATION L3 768 S L2 AND ADP L448 S L3 AND COMPOSITION L5 0 S L4 AND DIPTHERIA L612 S L4 AND TREATMENT L7 0 S L5 AND DIPHTHERIA L81 S L6 AND DIPHTHERIA L9 6 S L6 AND PERTUSSIS L10 1 S L6 AND TETANUS L110 S L6 AND INFECTION L12 1 S L4 AND ENTEROTOXIN L13 1 S L4 AND PROANTHOCYANIDIN L14 1 S L3 AND PROANTHOCYANIDIN

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             21 RIBOSYLATIONS
           3881 RIBOSYLATION
                  (RIBOSYLATION OR RIBOSYLATIONS)
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             37 ADPS
         30736 ADP
                  (ADP OR ADPS)
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           146 PROANTHOCYANIDIN
           141 PROANTHOCYANIDINS
           201 PROANTHOCYANIDIN
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L19
             0 L18 AND PROANTHOCYANIDIN
=> s 117 and proanthocyanidin
           146 PROANTHOCYANIDIN
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L20
             0 L17 AND PROANTHOCYANIDIN
=> s proanthocyanidin
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141 PROANTHOCYANIDINS
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=> s 121 and ADP
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L23
             1 L21 AND ADP
=> dis 123 bib abs
L23 ANSWER 1 OF 1
                       MEDLINE
ΑN
     2001526606
                    MEDLINE
DN
     21228268
              PubMed ID: 11330834
     Differential effects of IH636 grape seed proanthocyanidin
TΤ
     extract and a DNA repair modulator 4-aminobenzamide on liver microsomal
     cytochrome 4502E1-dependent aniline hydroxylation.
     Ray S D; Parikh H; Hickey E; Bagchi M; Bagchi D
ΑU
     Department of Pharmacology, Toxicology and Medicinal Chemistry, Arnold &
CS
     Marie Schwartz College of Pharmacy and Health Sciences, Long Island
     University, Brooklyn, New York 11201, USA.. sray@liu.edu
     MOLECULAR AND CELLULAR BIOCHEMISTRY, (2001 Feb) 218 (1-2) 27-33.
SO
     Journal code: NGU; 0364456. ISSN: 0300-8177.
CY
     Netherlands
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EΜ
     200109
    Entered STN: 20011001
ED
    Last Updated on STN: 20011001
    Entered Medline: 20010927
    Previous studies from our laboratories have linked the protective
AΒ
    abilities of IH636 grape seed proanthocyanidin extract (GSPE)
    with inactivation of anti-apoptotic gene bcl-XL, and modification of
    several other critical molecular targets such as DNA-damage/DNA-repair,
    lipid peroxidation and intracellular Ca2+ homeostasis. Especially, GSPE
    provided dramatic protection against acetaminophen (APAP)-induced
    hepatotoxicity, significantly increased bcl-XL expression in the liver,
    and antagonized both necrotic and apoptotic deaths of liver cells in vivo.
    However, it was not clear from this study whether anti-apoptogenic and
    anti-necrotic effects of GSPE were: (i) due to its interference with
    endonuclease activity, (ii) due to its antioxidant effect, or, (iii) due
    to its ability to inhibit microsomal drug metabolizing enzyme(s), such as
    CYP-4502E1. Since CYP-4502E1 primarily metabolizes acetaminophen in mice
    and rats, this study specifically focused on CYP-4502E1's catalytic
    activity in vitro. Overall this investigation compared the in vitro
    aniline hydroxylation patterns of: (i) in vivo GSPE-exposed and unexposed
    (control) mouse liver microsomes, (ii) induced (1% acetone in drinking
    water for 3 days) and uninduced rat liver microsomes in the presence and
    absence of GSPE in vitro, and (iii) control rat liver microsomes in the
    presence of an anti-APAP agent 4-aminobenzamide (4-AB) in vitro. For the
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in vivo assessment, male $B\acute{e}C3F1$ mice were fed GSPE diet (ADI 100 mg/kg body wt) for 4 weeks, and liver microsomes were isolated from both control

and GSPE-fed mice for aniline hydroxylation, a specific marker of CYP-4502E1 activity. Data show that hydroxylation was 40% less in microsomes from GSPE-exposed livers compared to control microsomes.

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Similarly, when rat liver microsomes were incubated with various concentrations of GSPE in vitro (100 and 250 microg/ml), aniline hydroxylation was inhibited to various degrees (uninduced: 40 and 60% and induced: 25 and 50%, respectively with 100 and 250 microg/ml). Influence of GSPE on hydroxylation patterns were compared with another hepatoprotective agent 4-aminobenzamide (4-AB), a well-known modulator of nuclear enzyme poly(ADP-ribose) polymerase, and the data shows that 4-AB did not alter aniline hydroxylation at all. Collectively, these results may suggest that GSPE has the ability to inhibit CYP-4502E1, and this is an additional cytoprotective attribute, in conjunction with its novel antioxidant and/or antiendonucleolytic potential.

=> dis hist

=> s 126 and treatment

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(FILE 'HOME' ENTERED AT 08:38:59 ON 16 MAY 2002)
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 L1
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 L3
             768 S L2 AND ADP
 L4
              48 S L3 AND COMPOSITION
 L5
              0 S L4 AND DIPTHERIA
 L6
              12 S L4 AND TREATMENT
 L7
              0 S L5 AND DIPHTHERIA
 L8
              1 S L6 AND DIPHTHERIA
 L9
              6 S L6 AND PERTUSSIS
 L10
              1 S L6 AND TETANUS
 L11
              0 S L6 AND INFECTION
 L12
               1 S L4 AND ENTEROTOXIN
 L13
               1 S L4 AND PROANTHOCYANIDIN
L14
               1 S L3 AND PROANTHOCYANIDIN
     FILE 'MEDLINE' ENTERED AT 08:52:37 ON 16 MAY 2002
         508108 S L1
L15
            979 S L2
L16
             954 S L3
L17
L18
              4 S L4
L19
              0 S L18 AND PROANTHOCYANIDIN
L20
              0 S L17 AND PROANTHOCYANIDIN
L21
            201 S PROANTHOCYANIDIN
L22
              0 S L21 AND RIBOSYLATION
L23
              1 S L21 AND ADP
=> s bacterial
        348875 BACTERIAL
             8 BACTERIALS
L24
        348876 BACTERIAL
                 (BACTERIAL OR BACTERIALS)
=> s 124 and infection
        393371 INFECTION
        450129 INFECTIONS
        674407 INFECTION
                 (INFECTION OR INFECTIONS)
L25
        120039 L24 AND INFECTION
=> s 125 and enterotoxin
          7398 ENTEROTOXIN
          8385 ENTEROTOXINS
          9792 ENTEROTOXIN
                 (ENTEROTOXIN OR ENTEROTOXINS)
L26
          1390 L25 AND ENTEROTOXIN
```

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1376847 TREATMENT 86934 TREATMENTS 1412705 TREATMENT

(TREATMENT OR TREATMENTS)

L27 100 L26 AND TREATMENT

=> s 127 and proanthocyanidin

146 PROANTHOCYANIDIN 141 PROANTHOCYANIDINS 201 PROANTHOCYANIDIN

(PROANTHOCYANIDIN OR PROANTHOCYANIDINS)

L28 0 L27 AND PROANTHOCYANIDIN

=>

---Logging off of STN---

=>

Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS FULL ESTIMATED COST	SINCE FILE ENTRY 4.38	TOTAL SESSION 61.27
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) CA SUBSCRIBER PRICE	SINCE FILE ENTRY 0.00	TOTAL SESSION -6.82

STN INTERNATIONAL LOGOFF AT 08:59:10 ON 16 MAY 2002